

WHAT IS CLAIMED IS:

- 1 1. A bioprosthetic heart valve comprising an acellular matrix and isolated myofibroblasts
2 wherein at least 60% of the total collagen produced by said myofibroblasts is type I
3 collagen.
- 1 2. The valve of claim 1, wherein said myofibroblasts produce at least 2-fold greater type I
2 collagen compared to type III collagen.
- 1 3. The valve of claim 1, wherein said myofibroblasts produce one or more extracellular
2 matrix components selected from the group consisting of fibronectin, elastin, and
3 glycosaminoglycan.
- 1 4. The valve of claim 3, wherein said glycosaminoglycan is chondroitin sulfate or
2 hyaluronic acid.
- 1 5. A valve comprising an acellular matrix and an isolated myofibroblast, wherein less than
2 25% of total collagen production by said myofibroblast is type III collagen.
- 1 6. The valve of claim 5, wherein less than 20% of total collagen production by said
2 myofibroblast is type III collagen.
- 1 7. The valve of claim 5, wherein less than 15% of total collagen production by said
2 myofibroblast is type III collagen.
- 1 8. The valve of claim 5, wherein said myofibroblast is derived from mammalian heart
2 leaflet interstitial tissue.
- 1 9. The valve of claim 5, wherein said myofibroblast is derived from a mammalian vascular
2 or dermal tissue.

- 1 10. The valve of claim 5, wherein said myofibroblast is derived from human heart leaflet
2 interstitial tissue.
- 1 11. A method of enhancing production of type I collagen by an isolated myofibroblast,
2 comprising culturing said myofibroblast under pulsatile flow conditions.
- 1 12. The method of claim 11, wherein said myofibroblast is cultured in the presence of basic
2 fibroblast growth factor.
- 1 13. The method of claim 11, wherein said myofibroblast is cultured in endothelial cell-
2 conditioned media.
- 1 14. The method of claim 11, wherein said myofibroblast is cultured in the presence of an
2 isolated endothelial cell.
- 1 15. A method of enhancing viability and contractile activity of myofibroblasts in vitro
2 comprising culturing said myofibroblast under pulsatile flow conditions.
- 1 16. The method of claim 15, wherein said myofibroblast is cultured in endothelial cell-
2 conditioned media.
- 1 17. The method of claim 15, wherein said myofibroblast is cultured in the presence of an
2 isolated endothelial cell.
- 1 18. The method of claim 15, wherein said myofibroblast is cultured in the presence of a
2 purified endothelial cell-derived growth factor, wherein said growth factor inhibits
3 apoptosis of said myofibroblast.

1 19. An isolated myofibroblast, wherein said myofibroblast is genetically altered to increase
2 type I collagen production relative to type III collagen production.

1 20. A bioprosthetic heart valve comprising the myofibroblast of claim 19.

1 21. A method of manufacturing an artificial heart valve, comprising

2 (a) providing an acellular matrix,

3 (b) seeding said matrix with isolated myofibroblasts; and

4 (c) culturing said myofibroblasts under pulsatile flow conditions.

1 22. The method of claim 21, wherein said myofibroblasts are derived from an intended
2 recipient from an intended recipient of said centrifugal heart valve.